SEQUENCE LISTING

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<110> Kaslow, David C. Tsuboi, Takafumi Torii, Motomi The Government of the United States of America as represented by the Secretary of the Department of Health and Human Services <120> Vaccines for Blocking Transmission of Plasmodium vivax <130> 015280-34210US <140> US 09/554,960 <141> 2003-02-12 <150> US 60/045,283 <151> 1997-05-01 <150> WO PCT/US98/25742 <151> 1998-12-04 <160> 24 <170> PatentIn Ver. 2.0 <210> 1 <211> 1066 <212> DNA <213> Plasmodium vivax <220> <221> CDS <222> (147)..(857) <223> Pvs28 <400> 1 tccactcctc tcttgttcca cactttatct ttgtttcccc ccattcggcc accaactgca 60 ttatacaaaa acgactcccc ctttgagata acacccaact gagctcgatt ccccctcccc 120 acttttgcgc ctcccccttg ttcaaa atg aat acc tac cac agc ttg ctg ttc Met Asn Thr Tyr His Ser Leu Leu Phe 1 ctt ctg gcc atc gtg ctt act gtt aag cac acc ttc gca aag gtc acc Leu Leu Ala Ile Val Leu Thr Val Lys His Thr Phe Ala Lys Val Thr 10 15 gcg gag acc caa tgc aaa aat ggc tat gta gtc caa atg agc aat cat Ala Glu Thr Gln Cys Lys Asn Gly Tyr Val Val Gln Met Ser Asn His 30 ttt gaa tgc aaa tgc aac gac ggg ttt gtt atg gca aat gaa aac act Phe Glu Cys Lys Cys Asn Asp Gly Phe Val Met Ala Asn Glu Asn Thr 45 tgc gag gaa aaa cgc gat tgc aca aat cca caa aat gta aat aaa aac Cys Glu Glu Lys Arg Asp Cys Thr Asn Pro Gln Asn Val Asn Lys Asn

. . . .

		tgt gca aac Cys Ala Asn 80						
		tgc ata tta Cys Ile Leu						
		tgt aac ggc Cys Asn Gly			Lys			
-	-	aat gtg aac Asn Val Asn 130	_					
		gaa tct aaa Glu Ser Lys 145						
•		tgt aag gca Cys Lys Ala 160						
~		gtt gcg aag Val Ala Lys			-			
		gga gag ggc Gly Glu Gly			Gly			
		gac aca gga Asp Thr Gly 210		_				
		ata cta ctt Ile Leu Leu 225						
tca tta gtg tagacgattc tacacacaca cacaaacata cacaagggga 8 Ser Leu Val 235								
gaagegtete acagagteag tteaagteat acgeacaaaa aaggaaagta catecagetg								
gtgaaagagc atttatgtgt gcagttatcc ttgggagaag caccctccac ccagttgcgt								
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<210> 2

<211> 236

<212> PRT

<213> Plasmodium vivax

<400> 2

Met Asn Thr Tyr His Ser Leu Leu Phe Leu Leu Ala Ile Val Leu Thr 1 5 10 15

Val Lys His Thr Phe Ala Lys Val Thr Ala Glu Thr Gln Cys Lys Asn 20 25 30

Gly Tyr Val Val Gln Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp $35 \hspace{1cm} 40 \hspace{1cm} 45$

Gly Phe Val Met Ala Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys
50 60

Thr Asn Pro Gln Asn Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys 65 70 75 80

Ala Asn Thr Arg Met Asn Asp Glu Glu Arg Ala Leu Arg Cys Gly Cys 85 90 95

Ile Leu Gly Tyr Thr Val Met Asn Glu Val Cys Thr Pro Asn Lys Cys
100 105 110

Asn Gly Val Leu Cys Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn 115 120 125

Val Asn Ser Thr Met Cys Ser Cys Asn Ile Gly Thr Thr Leu Asp Glu 130 135 140

Ser Lys Lys Cys Gly Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys 145 150 155 160

Lys Ala Asn Glu Glu Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val 165 170 175

Ala Lys Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly 180 185 190

Glu Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly Asp 195 200 205

Thr Gly Ala Ala Tyr Ser Leu Met Asn Gly Ser Ala Val Ile Ser Ile 210 215 220

Leu Leu Val Phe Ala Phe Phe Met Met Ser Leu Val 225 230 235

<210> 3

<211> 995

<212> DNA

<213> Plasmodium vivax

<220>

<221> CDS

<222> (255)..(914)

<223> Pvs25

<400> 3

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-

tccacttagc caaa	atg aac tc Met Asn Se 1								
gtc caa att gcg Val Gln Ile Ala 15									
ata tgc aaa aat Ile Cys Lys Asn 30		ı Val Gln							
atg tgt aac gaa Met Cys Asn Glu 45	ggg ctg gt Gly Leu Va 50	g cac ctt l His Leu	tcc gaa Ser Glu 55	a'at aca Asn Thr	tgt gaa Cys Glu	gaa 434 Glu 60			
aaa aat gaa tgc Lys Asn Glu Cys									
ggc cag tgt ata Gly Gln Cys Ile 80									
tgt ggt tgc att Cys Gly Cys Ile 95									
gat gta tgt caa Asp Val Cys Gln 110		n Cys Gly				_			
gag tac ctc tcg Glu Tyr Leu Ser 125									
aaa gtc ccc aat Lys Val Pro Asn			_			_			
act gct tgt caa Thr Ala Cys Gln 160	Leu Lys Cy								
gtt gaa gga gtt Val Glu Gly Val 175									
aaa gag aaa aat Lys Glu Lys Asn 190		ı Ser Tyr							
tac tcc ctc ttc Tyr Ser Leu Phe 205						911			
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<210> 4
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<211> 219

<212> PRT

<213> Plasmodium vivax

<400>4

Met Asn Ser Tyr Tyr Ser Leu Phe Val Phe Phe Leu Val Gln Ile Ala 1 5 10 15

Leu Lys Tyr Ser Lys Ala Ala Val Thr Val Asp Thr Ile Cys Lys Asn 20 25 30

Gly Gln Leu Val Gln Met Ser Asn His Phe Lys Cys Met Cys Asn Glu $35 \hspace{1cm} 40 \hspace{1cm} 45$

Gly Leu Val His Leu Ser Glu Asn Thr Cys Glu Glu Lys Asn Glu Cys
50 60

Lys Lys Glu Thr Leu Gly Lys Ala Cys Gly Glu Phe Gly Gln Cys Ile
65 70 75 80

Glu Asn Pro Asp Pro Ala Gln Val Asn Met Tyr Lys Cys Gly Cys Ile 85 90 95

Glu Gly Tyr Thr Leu Lys Glu Asp Thr Cys Val Leu Asp Val Cys Gln
100 105 110

Tyr Lys Asn Cys Gly Glu Ser Gly Glu Cys Ile Val Glu Tyr Leu Ser 115 120 125

Glu Ile Gln Ser Ala Gly Cys Ser Cys Ala Ile Gly Lys Val Pro Asn 130 135 140

Pro Glu Asp Glu Lys Lys Cys Thr Lys Thr Gly Glu Thr Ala Cys Gln 145 150 155 160

Leu Lys Cys Asn Thr Asp Asn Glu Val Cys Lys Asn Val Glu Gly Val
165 170 175

Tyr Lys Cys Gln Cys Met Glu Gly Phe Thr Phe Asp Lys Glu Lys Asn 180 185 190

Val Cys Leu Ser Tyr Ser Val Phe Asn Ile Leu Asn Tyr Ser Leu Phe 195 200 205

Phe Ile Ile Leu Leu Val Leu Ser Tyr Val Ile 210 215

<210> 5

<211> 377

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs25-Pvs28
 fusion protein

<400> 5

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1 5 10 15

- Ser Asn His Phe Lys Cys Met Cys Asn Glu Gly Leu Val His Leu Ser 20 25 30

 Glu Asn Thr Cys Glu Glu Lys Asn Glu Cys Lys Lys Glu Thr Leu Gly
- Lys Ala Cys Gly Glu Phe Gly Gln Cys Ile Glu Asn Pro Asp Pro Ala 50 55 60
- Gln Val Asn Met Tyr Lys Cys Gly Cys Ile Glu Gly Tyr Thr Leu Lys 65 70 75 80
- Glu Asp Thr Cys Val Leu Asp Val Cys Gln Tyr Lys Asn Cys Gly Glu 85 90 95
- Ser Gly Glu Cys Ile Val Glu Tyr Leu Ser Glu Ile Gln Ser Ala Gly
 100 105 110
- Cys Ser Cys Ala Ile Gly Lys Val Pro Asn Pro Glu Asp Glu Lys Lys 115 120 125
- Cys Thr Lys Thr Gly Glu Thr Ala Cys Gln Leu Lys Cys Asn Thr Asp 130 140
- Asn Glu Val Cys Lys Asn Val Glu Gly Val Tyr Lys Cys Gln Cys Met 145 150 155 160
- Glu Gly Phe Thr Phe Asp Lys Glu Lys Asn Val Cys Leu Ser Gly Gly 165 170 175
- Gly Pro Gly Gly Gly Ala Lys Val Thr Ala Glu Thr Gln Cys Lys Asn 180 185 190
- Gly Tyr Val Val Gln Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp 195 200 205
- Gly Phe Val Met Ala Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys 210 215 220
- Thr Asn Pro Gln Asn Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys 225 230 235 240
- Ala Asn Thr Arg Met Asn Asp Glu Glu Arg Ala Leu Arg Cys Gly Cys 245 250 255
- Ile Leu Gly Tyr Thr Val Met Asn Glu Val Cys Thr Pro Asn Lys Cys
 260 265 270
- Asn Gly Val Leu Cys Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn 275 280 285
- Val Asn Ser Thr Met Cys Ser Cys Asn Ile Gly Thr Thr Leu Asp Glu 290 295 300
- Ser Lys Lys Cys Gly Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys 305 310 315 320
- Lys Ala Asn Glu Glu Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val 325 330 335

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Ala Lys Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly
                                345
Glu Gly Ser Gly Gly Glu Gly Ser Gly Glu Gly Ser Gly Asp
Thr Gly Ala Ala Tyr Ser Leu Met Asn
<210> 6
<211> 21
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence:sense primer
<400> 6
                                                                   21
ggwtttytrr ytcaratgag t
<210> 7
<211> 17
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: vector-specific
     M13 universal primer
<400> 7
                                                                   17
gtaaaacgac ggccagt
<210> 8
<211> 24
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence:internal
     degenerate sense primer
<400> 8
                                                                   24
tcaratgagt rrycatttdg aatg
<210> 9
<211> 25
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:PCR-sense
      splinkerette #1 primer
<400> 9
                                                                   25
cgaatcgtaa ccgttcgtac gagaa
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<210> 10
<211> 20
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence:antisense Pvs25
      specific primer
ggacaagcag gatgataaag
                                                                    20
<210> 11
<211> 25
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:nested PCR
      sense splinkerette #2 internal primer
<400> 11
                                                                    25
tcgtaccaga atcgctgtcc tctcc
<210> 12
<211> 21
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:anti-sense
      Pvs25 specific internal primer
<400> 12
                                                                    21
agcacacaag tgtcttcctt c
<210> 13
<211> 18
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence:gene specific
      PCR sense primer
<400> 13
                                                                    18
actttcgttt cacagcac
<210> 14
<211> 20
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:gene specific
      PCR anti-sense primer
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20

<400> 14

aaaggacaag caggatgata

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<210> 15
<211> 7
<212> PRT
<213> Artificial Sequence
<223> Description of Artificial Sequence:flexible linker
<400> 15
Gly Gly Gly Pro Gly Gly Gly
<210> 16
<211> 186
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Pvs25 fusion
<400> 16
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Gln Leu Val Gln Met Ser Asn His Phe Lys Cys Met Cys Asn Glu Gly
Leu Val His Leu Ser Glu Asn Thr Cys Glu Glu Lys Asn Glu Cys Lys
Lys Glu Thr Leu Gly Lys Ala Cys Gly Glu Phe Gly Gln Cys Ile Glu
     50
Asn Pro Asp Pro Ala Gln Val Asn Met Tyr Lys Cys Gly Cys Ile Glu
Gly Tyr Thr Leu Lys Glu Asp Thr Cys Val Leu Asp Val Cys Gln Tyr
                 8.5
Lys Asn Cys Gly Glu Ser Gly Glu Cys Ile Val Glu Tyr Leu Ser Glu
Ile Gln Ser Ala Gly Cys Ser Cys Ala Ile Gly Lys Val Pro Glu Pro
                                                 125
        115
                            120
Glu Asp Glu Lys Lys Cys Thr Lys Thr Gly Glu Thr Ala Cys Gln Leu
                        135
Lys Cys Asn Thr Asp Asn Glu Val Cys Lys Asn Val Glu Gly Val Tyr
                    150
                                        155
                                                             160
Lys Cys Gln Cys Met Glu Gly Phe Thr Phe Cys Lys Glu Lys Asn Val
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170

175

165

Cys Leu Gly Pro His His His His His His 180 185

<210> 17

<211> 205

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs28 fusion protein

<400> 17

Glu Ala Glu Ala Ser Lys Val Thr Ala Glu Thr Gln Cys Lys Asn Gly
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Tyr Val Val Gln Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp Gly 20 25 30

Phe Val Leu Ala Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys Thr 35 40 45

Asn Pro Gln Asn Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys Ala
50 55 60

Asn Thr Arg Met Asn Asn Glu Glu Arg Ala Leu Arg Cys Gly Cys Ile 65 70 75 80

Leu Gly Tyr Thr Val Met Asn Glu Val Cys Thr Pro Tyr Lys Cys Asn 85 90 95

Gly Val Leu Cys Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn Val

Asn Ser Thr Met Cys Ser Cys Asn Ile Gly Ser Thr Leu Asp Glu Ser 115 120 125

Lys Lys Cys Gly Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys Lys 130 135

Ala Asn Glu Glu Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val Ala 145 150 155 160

Lys Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Glu 165 170 175

Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly Asp Thr 180 185 190

Gly Ala Ala Tyr Ser Gly Pro His His His His His 195 200 205

<210> 18

<211> 205

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs28Q130
 fusion protein

<400> 18

Glu Ala Glu Ala Ser Lys Val Thr Ala Glu Thr Gln Cys Lys Asn Gly
1 5 10 15

Tyr Val Val Gln Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp Gly 20 25 30

Phe Val Leu Ala Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys Thr 35 40 45

Asn Pro Gln Asn Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys Ala
50 55 60

Asn Thr Arg Met Asn Asn Glu Glu Arg Ala Leu Arg Cys Gly Cys Ile 65 70 75 80

Leu Gly Tyr Thr Val Met Asn Glu Val Cys Thr Pro Tyr Lys Cys Asn 85 90 95

Gly Val Leu Cys Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn Val 100 105 110

Gln Ser Thr Met Cys Ser Cys Asn Ile Gly Ser Thr Leu Asp Glu Ser 115 120 125

Lys Lys Cys Gly Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys Lys 130 135 140

Ala Asn Glu Glu Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val Ala 145 150 155 160

Lys Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Glu Glu Gly Glu 165 170 175

Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly Asp Thr 180 185 190

Gly Ala Ala Tyr Ser Gly Pro His His His His His His 195 200 205

<210> 19

<211> 169

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs28NCR fusion protein

<400> 19

Glu Ala Glu Ala Ser Lys Val Thr Ala Glu Thr Gln Cys Lys Asn Gly
1 5 10 15

Tyr Val Val Gln Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp Gly
20 25 30

Phe Val Leu Ala Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys Thr 35 40 45

Asn Pro Gln Asn Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys Ala
50 55 60

Asn Thr Arg Met Asn Asn Glu Glu Arg Ala Leu Arg Cys Gly Cys Ile
65 70 75 80

Leu Gly Tyr Thr Val Met Asn Glu Val Cys Thr Pro Tyr Lys Cys Asn 85 90 95

Gly Val Leu Cys Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn Val 100 105 110

Asn Ser Thr Met Cys Ser Cys Asn Ile Gly Ser Thr Leu Asp Glu Ser 115 120 125

Lys Lys Cys Gly Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys Lys 130 135

Ala Asn Glu Glu Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val Ala 145 150 155 160

Lys Gly Pro His His His His His His 165

<210> 20

<211> 174

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs25 domain of Pvs25-Pvs28 fusion protein

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Ser Asn His Phe Lys Cys Met Cys Asn Glu Gly Leu Val His Leu Ser 20 25 30

Glu Asn Thr Cys Glu Glu Lys Asn Glu Cys Lys Lys Glu Thr Leu Gly 35 40 45

Lys Ala Cys Gly Glu Phe Gly Gln Cys Ile Glu Asn Pro Asp Pro Ala 50 55 60

Gln Val Asn Met Tyr Lys Cys Gly Cys Ile Glu Gly Tyr Thr Leu Lys 65 70 75 80

Glu Asp Thr Cys Val Leu Asp Val Cys Gln Tyr Lys Asn Cys Gly Glu 85 90 95

Ser Gly Glu Cys Ile Val Glu Tyr Leu Ser Glu Ile Gln Ser Ala Gly
100 105 110

Cys Ser Cys Ala Ile Gly Lys Val Pro Asn Pro Glu Asp Glu Lys Lys 115 120 125 Cys Thr Lys Thr Gly Glu Thr Ala Cys Gln Leu Lys Cys Asn Thr Asp 130 135 140

Asn Glu Val Cys Lys Asn Val Glu Gly Val Tyr Lys Cys Gln Cys Met 145 150 155 160

Glu Gly Phe Thr Phe Asp Lys Glu Lys Asn Val Cys Leu Ser 165 170

<210> 21

<211> 196

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Pvs28 domain of Pvs25-Pvs28 fusion protein

<400> 21

Ala Lys Val Thr Ala Glu Thr Gln Cys Lys Asn Gly Tyr Val Val Gln 1 5 10 15

Met Ser Asn His Phe Glu Cys Lys Cys Asn Asp Gly Phe Val Met Ala 20 25 30

Asn Glu Asn Thr Cys Glu Glu Lys Arg Asp Cys Thr Asn Pro Gln Asn 35 40 45

Val Asn Lys Asn Cys Gly Asp Tyr Ala Val Cys Ala Asn Thr Arg Met 50 55 60

Asn Asp Glu Glu Arg Ala Leu Arg Cys Gly Cys Ile Leu Gly Tyr Thr 65 70 75 80

Val Met Asn Glu Val Cys Thr Pro Asn Lys Cys Asn Gly Val Leu Cys 85 90 95

Gly Lys Gly Lys Cys Ile Leu Asp Pro Ala Asn Val Asn Ser Thr Met
100 105 110

Cys Ser Cys Asn Ile Gly Thr Thr Leu Asp Glu Ser Lys Lys Cys Gly
115 120 125

Lys Pro Gly Lys Thr Glu Cys Thr Leu Lys Cys Lys Ala Asn Glu Glu 130 135 140

Cys Lys Glu Thr Gln Asn Tyr Tyr Lys Cys Val Ala Lys Gly Ser Gly 145 150 155 160

Gly Glu Gly Ser Gly Glu Gly Ser Gly Glu Gly Ser Gly Gly Glu 175

Glu Gly Ser Gly Glu Gly Ser Gly Gly Asp Thr Gly Ala Ala Tyr 180 185 190

Ser Leu Met Asn 195

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<210> 22
<211> 4
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<213> Artificial Sequence
<223> Description of Artificial Sequence: sequence added
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<400> 22
Glu Ala Glu Ala
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<210> 23
<211> 9
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: sequence added
      to enhance cleavage of alpha factor leader
<400> 23
Glu Ala Glu Ala Glu Ala Lys
                  5
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<210> 24
<211> 6
<212> PRT
<213> Artificial Sequence
<223> Description of Artificial Sequence:polyhistidine
      tag
<400> 24
His His His His His
                  5
  1
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